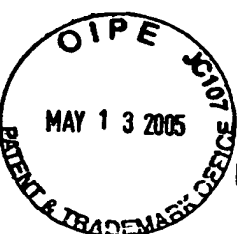


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JC06 Rec'd PCT/PTO 13 MAY 2005
PATENT



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

IN THE APPLICATION OF:

STEVEN GUTTERIDGE ET. AL.

CASE NO.: BB1533USPCT

APPLICATION NO.: 10/528611

CONFIRMATION NO.: UNKNOWN

GROUP ART UNIT: UNKNOWN

EXAMINER: UNASSIGNED

FILED: March 21, 2005

FOR: ISOLATION AND USE OF RYANODINE RECEPTORS

INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

Sir:

In compliance with 37 C.F.R. §§1.97 and 1.98, Applicants bring to the attention of the U.S. Patent and Trademark Office the information listed on the enclosed PTO/SB/08A and/or PTO/SB/08B forms. A copy of the information, if required, is also enclosed. Consideration of the information is requested under 37 C.F.R. § 1.56 and this information is submitted in accord with the provisions of §1.97(b): within three months of filing the national application under 37 C.F.R. §1.53(d), or entry into the national stage under 37 C.F.R. §1.491, or before the mailing of a first Office Action on the merits, or before the mailing of a first Office Action on the merits after filing a Request for Continued Examination under 37 C.F.R. §1.114.

If this Information Disclosure Statement is filed under §1.97(b) but the mailing date hereof is after the mailing of a first Office Action on the merits, the PTO is authorized to charge the fee set forth in 37 C.F.R. §1.17(p) to Deposit Account No. 04-1928 (E. I. du Pont de Nemours and Company) in order to

complete the requirements for consideration of this Information Disclosure Statement.

Please note we are also enclosing a copy of the International Search Report along with copies of those references cited therein.

Respectfully submitted,



JONATHON NARITA

Agent for Applicants

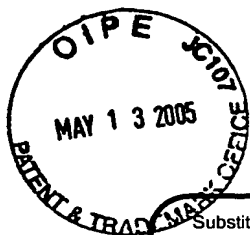
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STATEMENT BY APPLICANT**

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Sheet 1 of 5

Complete if Known

Application Number	10/528611
Filing Date	March 21, 2005
First Named Inventor	STEVEN GUTTERIDGE ET. AL.
Group Art Unit	UNKNOWN
Examiner Name	UNASSIGNED
Attorney Docket Number	BB1533USPCT

OTHER PRIOR ART -- NON PATENT LITERATURE DOCUMENTS

Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
	1	CHRISTOPHER H. GEORGE ET AL., Ryanodine Receptor Mutations Associated With Stress-Induced Ventricular Tachycardia Mediate Increased Calcium Release in Stimulated Cardiomyocytes, Circ. Res. 93:531-540, 2003	<input type="checkbox"/>
	2	KINYA OTSU ET AL., Chromosome Mapping of Five Human Cardiac and Skeletal Muscle Sarcoplasmic Reticulum Protein Genes, Genomics, 17:507-509, 1993	<input type="checkbox"/>
	3	GIUSEPPE GIANNINI ET AL., The Ryanodine Receptor/Calcium Channel Genes are Widely and Differentially Expressed in Murine Brain and Peripheral Tissues, The Journal of Cell Biology, 128(5):893-904, 1995	<input type="checkbox"/>
	4	DAWEI JIANG ET AL., Enhanced Basal Activity of a Cardiac Ca ²⁺ Release channel (Ryanodine Receptor) Mutant Associated with Ventricular Tachycardia and Sudden Death, Circulation Research, 91:218-225, 2002	<input type="checkbox"/>
	5	XUEHONG XU ET AL., Molecular Cloning of cDNA Encoding a Drosophila Ryanodine Receptor and Functional Studies of the Carboxyl-Terminal Calcium Release Channel, Biophysical Journal, 78:1270-1281, 2000	<input type="checkbox"/>
	6	HIROSHI TAKESHIMA ET AL., Ca ²⁺ -induced Ca ²⁺ release in myocytes from dyspedic mice lacking the type-1 ryanodine receptor, The EMBO Journal 14(13):2999-3006, 1995	<input type="checkbox"/>
	7	STEVEN O. MARX ET AL., PKA Phosphorylation Dissociates FKBP12.6 from the Calcium Release Channel (Ryanodine Receptor): Defective Regulation in Failing Hearts, Cell, Vol. 101:365-376, 2000	<input type="checkbox"/>
	8	ANDREW J. DINSMORE ET AL., Characterisation of Antibody Models of the Ryanodine Receptor for Use in High-Throughput Screening, Pestic Sci., Vol. 54:345-352, 1998	<input type="checkbox"/>
	9	TOSHIKI IMAGAWA ET AL., Expression of Ca ²⁺ -induced Ca ²⁺ Release Channel Activity from Cardiac Ryanodine Receptor cDNA in Chinese Hamster Ovary Cells, J. Biochem., Vol. 112:508-513, 1992	<input type="checkbox"/>
	10	BARBARA BAUCE ET AL., Screening for Ryanodine Receptor Type 2 Mutations in Families with Effort-Induced Polymorphic Ventricular Arrhythmias and Sudden Death, J. of Amer. Coll. of Card., Vol. 40(2):341-349, 2002	<input type="checkbox"/>
	11	GIAN ANTONIO ET AL., Genetics of arrhythmogenic right ventricular cardiomyopathy, Current Opinion in Cardiology, Vol. 17:218-221, 2002	<input type="checkbox"/>

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Considered

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Complete if Known

Application Number	10/528611
Filing Date	March 21, 2005
First Named Inventor	STEVEN GUTTERIDGE ET. AL.
Group Art Unit	UNKNOWN
Examiner Name	UNASSIGNED
Attorney Docket Number	BB1533USPCT

OTHER PRIOR ART -- NON PATENT LITERATURE DOCUMENTS

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	12	MIEKO SHIWA ET AL., Molecular Cloning and characterization of ryanodine receptor from unfertilized sea urchin eggs, Am. J. Physiol. Reg. Integrative Comp., Vol. 282:R727-R737, 2002	<input type="checkbox"/>
	13	YASUO OGAWA ET AL., Ryanodine Receptor Isoforms in Excitation-Contraction Coupling, Adv. Biophys., Vol. 26:27-64, 1999	<input type="checkbox"/>
	14	G. LEES ET AL., Cell Culture Approaches to Invertebrae Neuroscience, Academic Press, New York, pp. 123-127, 1988	<input type="checkbox"/>
	15	MANJUNATHA B. BHAT ET AL., Functional Calcium release Channel Formed by the Carboxyl-Terminal Portion of Ryanodine Receptor, Biophysical J., Vol. 73:1329-1336, 1997	<input type="checkbox"/>
	16	NATASCIA TISO ET AL., The binding of the RyR2 calcium channel to its gating protein FKBP12.6 is oppositely affected by ARVD2 and VTSIP mutations, Biochem. & Biophys. Res. Comm., Vol. 229:594-598, 2002	<input type="checkbox"/>
	17	ISAAC N. PESSAH ET AL., Calcium-Ryanodine Receptor Complex, The J. of Biol. Chem., Vol. 261(19):8643-8648, 1986	<input type="checkbox"/>
	18	ELISABETH LEHMBERG ET AL., Similarity of Insect and Mammalian Ryanodine Binding Sites, Pesticide Biochem. & Phys., Vol. 48:145-152, 1994	<input type="checkbox"/>
	19	HIROSHI TAKESHIMA ET AL., Isolation and characterization of a gene for a ryanodine receptor/calcium release channel in Drosophila melanogaster, FEBS Letters, Vol. 337:81-87, 1994	<input type="checkbox"/>
	20	NATIONAL CENTER FOR BIOTECHNOLOGY INFORMATION GENERAL IDENTIFIER NO. 456161, ACC. NO. D17389, 3/25/1999, H. TAKESHIMA ET AL., Isolation and characterization of a gene for a ryanodine receptor/calcium release channel in Drosophila melanogaster	<input type="checkbox"/>
	21	NATIONAL CENTER FOR BIOTECHNOLOGY INFORMATION GENERAL IDENTIFIER NO. 1871446, ACC. NO. D45899, 12/25/2002, Y. SAKUBE ET AL., An abnormal ketamine response in mutants defective in the ryanodine receptor gene ryr-1 (unc-68) of Caenorhabditis elegans	<input type="checkbox"/>
	22	YASUJI SAKUBE ET AL., An Abnormal ketamine Response in Mutants Defective in the Ryanodine Receptor Gene ryr-1 (unc-68) of Caenorhabditis elegans, J. Mol. Biol., Vol. 267:849-864, 1997	<input type="checkbox"/>

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**INFORMATION DISCLOSURE
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Sheet 3 of 5

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Application Number	10/528611
Filing Date	March 21, 2005
First Named Inventor	STEVEN GUTTERIDGE ET. AL.
Group Art Unit	UNKNOWN
Examiner Name	UNASSIGNED
Attorney Docket Number	BB1533USPCT

OTHER PRIOR ART -- NON PATENT LITERATURE DOCUMENTS

Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
	23	ELENA PUENTE ET AL., Identification of a polymorphic ryanodine receptor gene from <i>Heliothis virescens</i> (Lepidoptera: Noctuidae), Insect Biochem. & Mol. Biol., Vol. 30:335-347, 2000	<input type="checkbox"/>
	24	NATIONAL CENTER FOR BIOTECHNOLOGY INFORMATION GENERAL IDENTIFIER NO. 17352471, ACC. NO. 476994, 12/10/2003, M. D. ADAMS ET AL., The genome sequence of <i>Drosophila Melanogaster</i>	<input type="checkbox"/>
	25	MARK D. ADAMS ET AL., The genome sequence of <i>Drosophila Melanogaster</i> , Vol. 287:2185-2195, 2000	<input type="checkbox"/>
	26	NATIONAL CENTER FOR BIOTECHNOLOGY INFORMATION GENERAL IDENTIFIER NO. 21301556, ACC. NO. EAA17301, 5/31/2002	<input type="checkbox"/>
	27	NATIONAL CENTER FOR BIOTECHNOLOGY INFORMATION GENERAL IDENTIFIER NO. 1871447, ACC. NO. BAA08309, 12/25/2002, Y. SAKUBE ET AL., An abnormal ketamine response in mutants defective in the ryanodine receptor gene <i>ryr-1</i> (unc-68) of <i>Caenorhabditis elegans</i>	<input type="checkbox"/>
	28	NATIONAL CENTER FOR BIOTECHNOLOGY INFORMATION GENERAL IDENTIFIER NO. 18656155, ACC. NO. BAB84714, 2/14/2002, M. SHIWA ET AL., Molecular cloning and characterization of ryanodine receptor from unfertilized sea urchin eggs	<input type="checkbox"/>
	29	NATIONAL CENTER FOR BIOTECHNOLOGY INFORMATION GENERAL IDENTIFIER NO. 13569850, ACC. NO. NP076357, 12/21/2003, H. MASUMIYA ET AL., The mouse sino-atrial node expresses both the type 2 and type 3 (Ca ²⁺) release channels/ryanodine receptors	<input type="checkbox"/>
	30	HARUKO MASUMIYA ET AL., The mouse sino-atrial node expresses both the type 2 and type 3 (Ca ²⁺) release channels/ryanodine receptors, Febs Letters, Vol. 553:141-144, 2003	<input type="checkbox"/>
	31	PATRICK MOST ET AL., Transgenic Overexpression of the Ca ²⁺ -binding Protein S100A1 in the Heart Leads to Increased in Vivo Myocardial Contractile Performance, J. of Biol. Chem., Vol. 278(36):33809-33817, 2003	<input type="checkbox"/>
	32	HUANG-TIAN YANG ET AL., The ryanodine receptor modulates the spontaneous beating rate of cardiomyocytes during development, PNAS, Vol. 99(14):9225-9230, 2002	<input type="checkbox"/>
	33	ANNE-VALERIE FAURE ET AL., Developmental expression of the calcium release channels during early neurogenesis of the mouse cerebral cortex, European J. of Neuroscience, Vol. 14:1613-1622, 2001	<input type="checkbox"/>

Examiner
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**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**

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Sheet 4 of 5

Complete if Known

Application Number	10/528611
Filing Date	March 21, 2005
First Named Inventor	STEVEN GUTTERIDGE ET. AL.
Group Art Unit	UNKNOWN
Examiner Name	UNASSIGNED
Attorney Docket Number	BB1533USPCT

OTHER PRIOR ART -- NON PATENT LITERATURE DOCUMENTS

Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
	34	MINGCAI ZHAO ET AL., Molecular identification of the Ryanodine Receptor Pore-forming Segment, J. of Biol. Chem., Vol. 274(37):25971-25974, 1999	<input type="checkbox"/>
	35	HIROSHI TAKESHIMA ET AL., Embryonic lethality and abnormal cardiac myocytes in mice lacking ryanodine receptor type 2, The EMBO J., Vol. 17(12):3309-3316, 1998	<input type="checkbox"/>
	36	NATIONAL CENTER FOR BIOTECHNOLOGY GENERAL IDENTIFIER NO. 1245376, ACC. NO. AAA93465, 4/2/1996, J. NAKAI ET AL., Primary structure and functional expression from cDNA of the cardiac ryanodine receptor/calcium release channel	<input type="checkbox"/>
	37	JUNICHI NAKAI ET AL., Primary structure and functional expression from cDNA of the cardiac ryanodine receptor/calcium release channel, FEBS, Vol. 271(1,2):169-177, 1990	<input type="checkbox"/>
	38	NATIONAL CENTER FOR BIOTECHNOLOGY INFORMATION GENERAL IDENTIFIER NO. 4506757, ACC. NO. NP001026, 12/23/2003, C.H. GEORGE ET AL., Ryanodine receptor mutations associated with stress-induced ventricular tachycardia mediate increased calcium release in stimulated cardiomyocytes	<input type="checkbox"/>
	39	CHRISTOPHER H. GEORGE ET AL., Ryanodine receptor mutations associated with stress-induced ventricular tachycardia mediate increased calcium release in stimulated cardiomyocytes, J. of Biol. Chem., Vol. 278(31):28856-28864, 2003	<input type="checkbox"/>
	40	JING ZHANG ET AL., Three-dimensional Localization of Divergent Region 3 of the Ryanodine Receptor to the Clamp-shaped Structures Adjacent to the FKBP Binding Sites, J. Biol. Chem., Vol. 278(16):14211-14218, 2003	<input type="checkbox"/>
	41	HARUKO MASUMIYA ET AL., Localization of the 12.6-kDa FK506-binding Protein (FKBP12.6) Binding Site to the NH2-terminal Domain of the Cardiac Ca2+ Release Channel (Ryanodine Receptor), J. Biol. Chem., Vol. 278(6):3786-3792, 2003	<input type="checkbox"/>
	42	JIEFEI TONG ET AL., Caffeine and Halothane Sensitivity of Intracellular Ca2+ Release is Altered by 15 Calcium Release Channel (Ryanodine Receptor) Mutations Associated with Malignant Hyperthermia and/or Central Core Disease, J. Biol. Chem., Vol. 272(42):26332-26339, 1997	<input type="checkbox"/>
	43	S. R. WAYNE CHEN ET AL., Antibodies as Probes for Ca2+ Activation Sites in the Ca2+ Release Channel (Ryanodine Receptor) of Rabbit Skeletal Muscle Sarcoplasmic Reticulum, J. Biol. Chem., Vol. 268(18):13414-13421, 1993	<input type="checkbox"/>
	44	CELETTA CALLWAY ET AL., Localization of the High and Low Affinity [3H]Ryanodine binding Sites on the Skeletal Muscle Ca2+ Release Channel, J. Biol. Chem., Vol. 269(22):15876-15884, 1994	<input type="checkbox"/>

Examiner Signature	Date Considered
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Sheet 5 of 5

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First Named Inventor	STEVEN GUTTERIDGE ET. AL.
Group Art Unit	UNKNOWN
Examiner Name	UNASSIGNED
Attorney Docket Number	BB1533USPCT

OTHER PRIOR ART -- NON PATENT LITERATURE DOCUMENTS

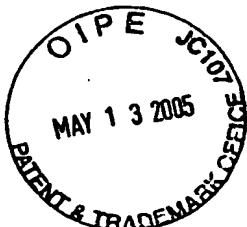
Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
	45	MELANIE SCHMITT ET AL., Binding Sites for Ca ²⁺ -Channel Effectors and Ryanodine in Periplaneta americana - Possible Targets for New Insecticides, Pestic Science, Vol. 48:375-385, 1996	<input type="checkbox"/>
	46	RICHARD E.A. TUNWELL ET AL., The human cardiac muscle ryanodine receptor-calcium release channel: identification, primary structure and topological analysis, Biochem. J. Vol. 318:477-487, 1996	<input type="checkbox"/>
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